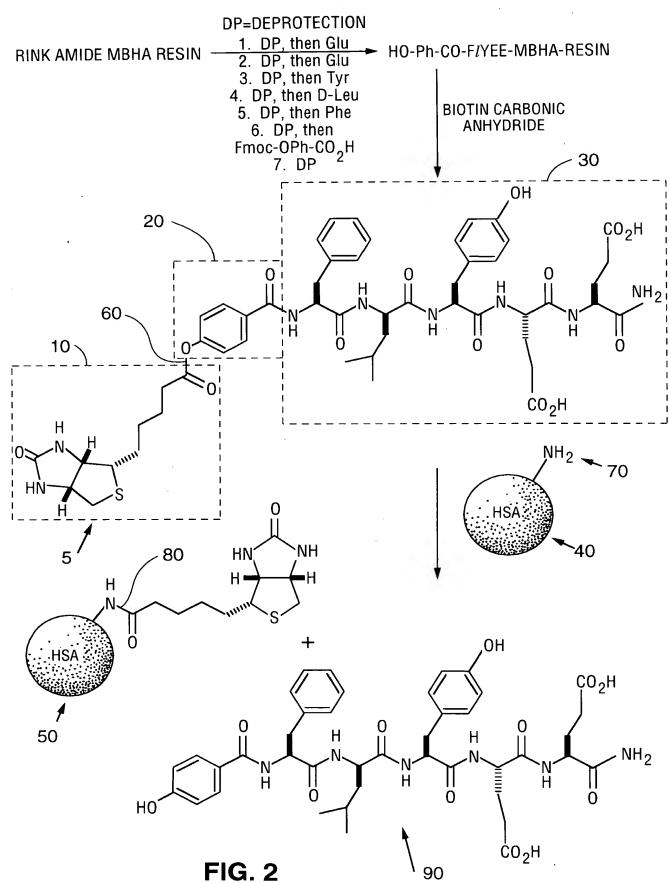
FIG. 1

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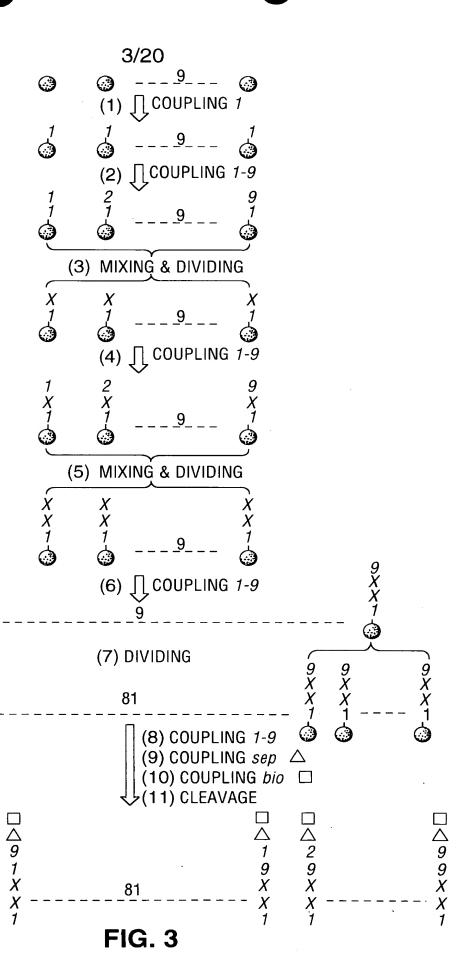
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APPROVED O.G. FIG. .

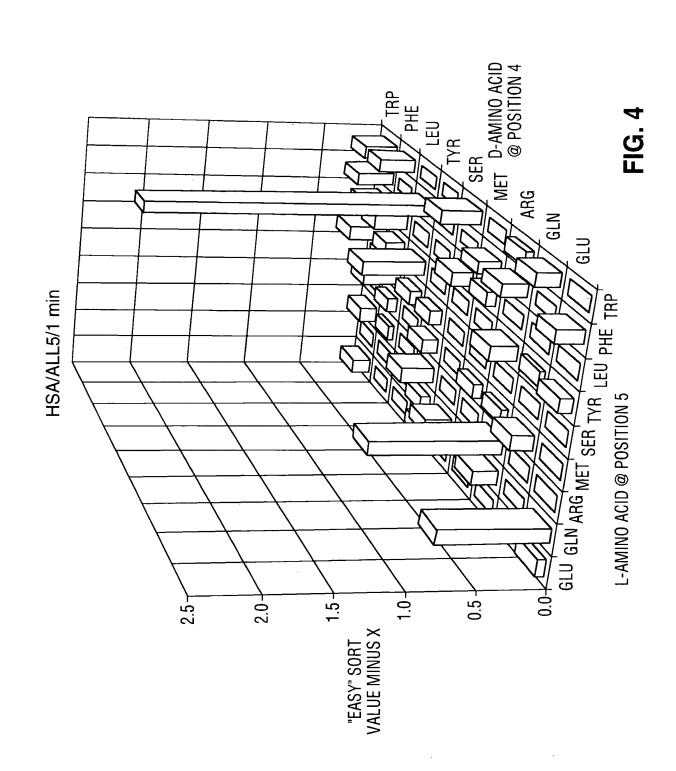
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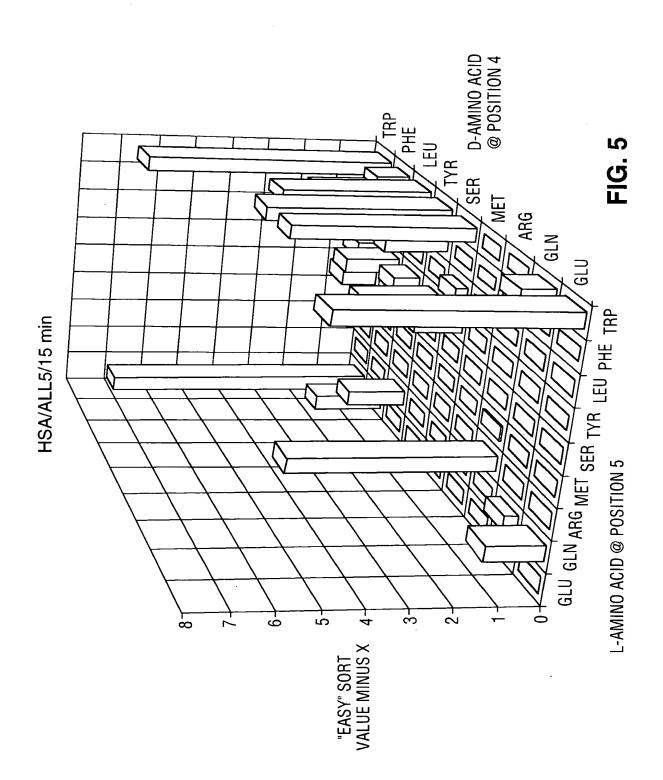
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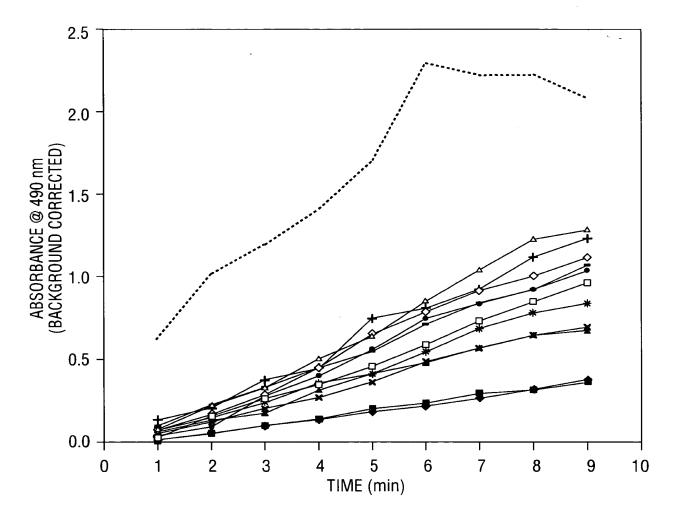




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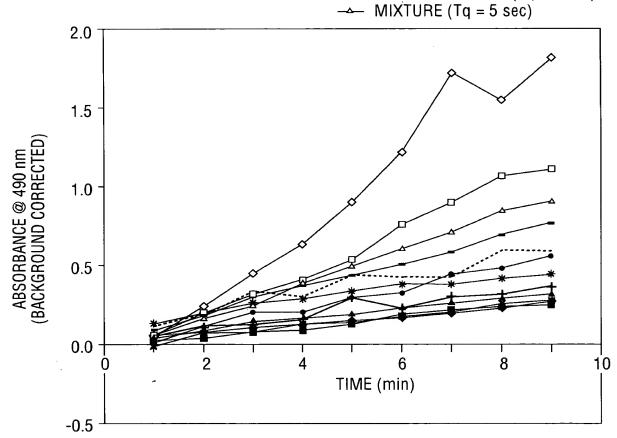
REACTION OF HSA WITH BIOTIN-S-(PHENYL)CONH-F- \underline{L} -O-X-NH2; QUENCH TIME = 5 sec.

FIG. 6A

APPROVEG	O.G. FIG.	
BY 4	CLASS	SUBCLASS
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→ S(phenyl)-CONH2 (Tq = 5 sec)
→ S(phenyl)-CONH2 (Tq = 5 sec)
→ x-W-E-E-X-E-NH2 (Tq = 5 sec)
→ x-W-E-Q-X-E-NH2 (Tq = 5 sec)
→ x-W-E-R-X-E-NH2 (Tq = 5 sec)
→ x-W-E-M-X-E-NH2 (Tq = 5 sec)
→ x-W-E-S-X-E-NH2 (Tq = 5 sec)
- x-W-E-Y-X-E-NH2 (Tq = 5 sec)
→ x-W-E-L-X-E-NH2 (Tq = 5 sec)
→ x-W-E-F-X-E-NH2 (Tq = 5 sec)
- x-W-E-W-X-E-NH2 (Tq = 5 sec)



REACTION OF HSA WITH BIOTIN-S-(PHENYL)p CONH-W- $\underline{\mathbf{E}}$ -O-X-NH2; QUENCH TIME = 5 SEC.

FIG. 6B

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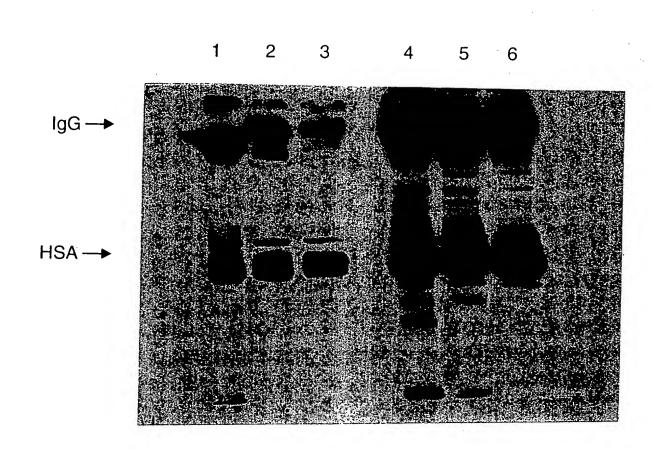
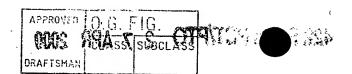
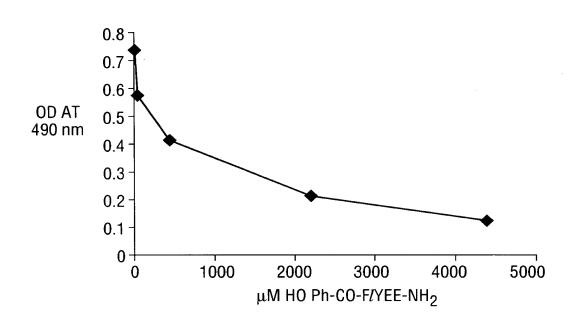


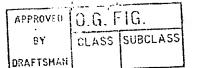
FIG. 7

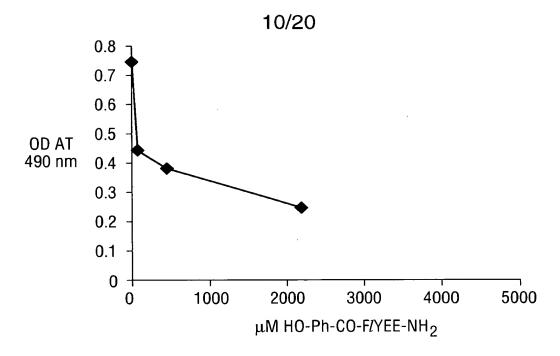




COMPETITION OF BIOTIN-OPh-CO-F l YEE-NH $_2$ (44 μ M) vs. HO-Ph-CO-F l YEE-NH $_2$

FIG. 8





"COMPETITION" TEST BETWEEN BIOTIN-OPh-FlYEE-NH $_2$ (44 μ M) AND HO-Ph-CO-FlYEE-NH $_2$

FIG. 9

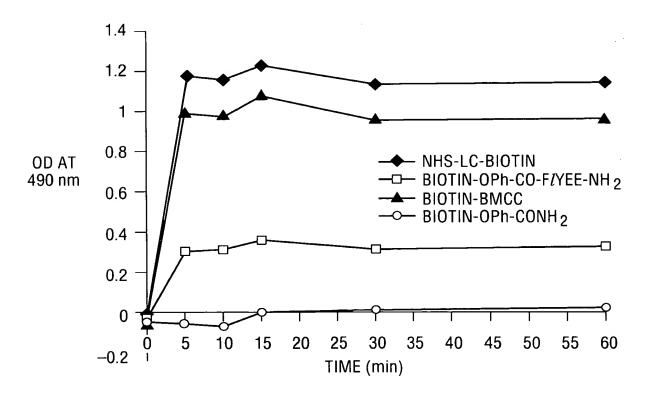
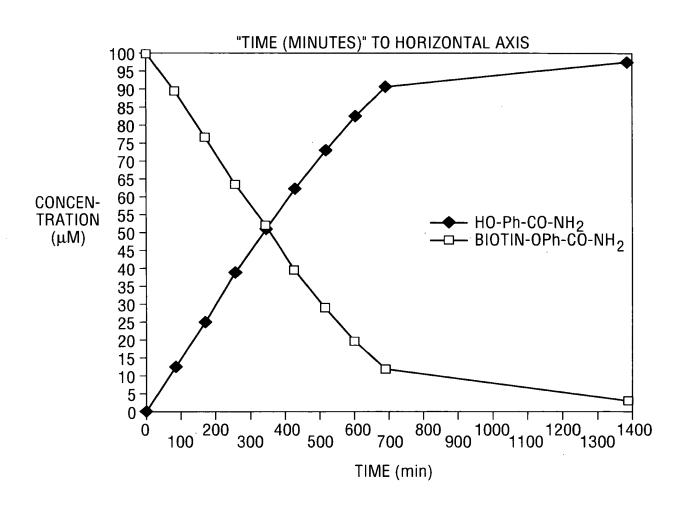


FIG. 10

APPROVED	o.g. Fig.		
54.	CLASS	SUBCLASS	
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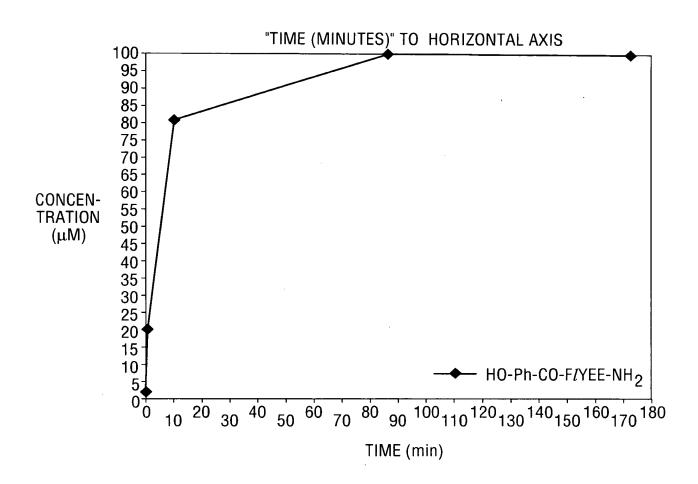


RATE OF REACTION OF 600 μM HSA WITH 100 μM BIOTIN-OPh-CONH₂ IN pH 7.4 PBS AT RT

FIG. 11

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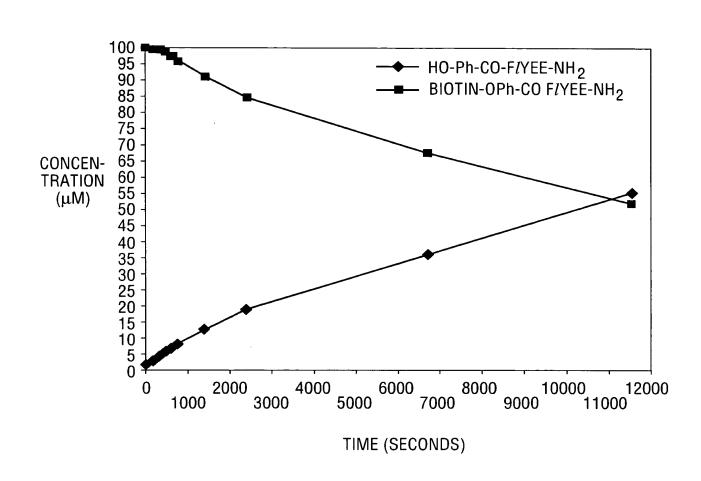
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RATE OF REACTION OF 100 μ M BIOTIN-OPh-CO-F*i*YEE-NH $_2$ IN COMMERCIAL HUMAN PLASMA

FIG. 12

APPROVEO	O.G. FIG.	
ey	CLASS	SUBCLASS
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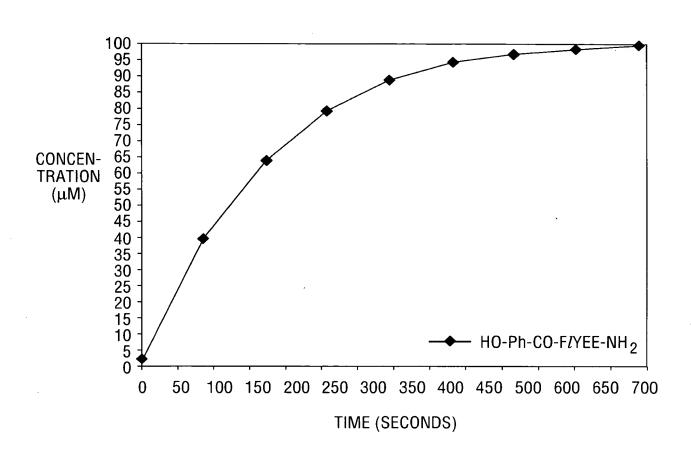
RATE OF HYDROLYSIS OF 100 μM BIOTIN-OPh-CO-F/YEE-NH $_2$ IN pH 7.4 PBS AT RT

FIG. 13

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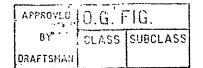
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RATE OF REACTION OF 100 μM BIOTIN-OPh-CO-FIYEE-NH $_2$ WITH 600 μM HSA AT RT

FIG. 14



"DEVICE A" FROM HSACONTROL(BOPhF/YEE)RCM-TD

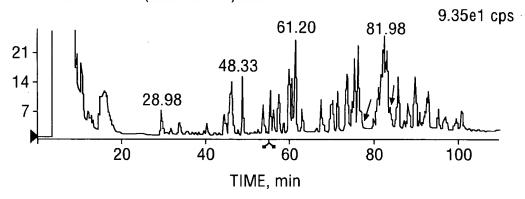


FIG. 15A

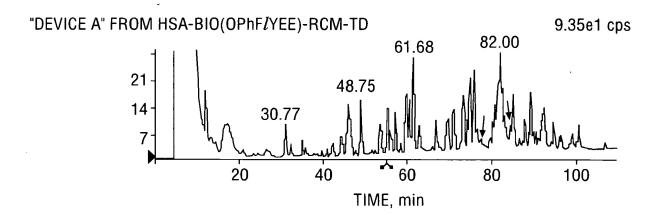


FIG. 15B

APPROVEG	O.G. FIG.		
BY	CLASS	SUBCLASS	
DRAFTSMAH			

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BIOTIN ISOBUTYLCARBONIC ANHYDRIDE

TEA/DMF 2. TFA/H₂ O/PHENOL

DP=DEPROTECT WITH 20% PIPERIDINE

DP, then Fmoc-Glu(OtBu)-OH DP, then Fmoc-Glu(OtBu)-OH

DP, then Fmoc-Tyr(tBu)-OH

DP, then Fmoc-D-Leu-OH

Fmoc-RINK AMIDE MBHA RESIN

DP=DEPROTECT WITH 20% PIPERIDINE

then Fmoc-D-Leu-OH

Fmoc-RINK AMIDE MBHA RESIN

SOBUTYLCARBONIC ANHYDRIDE TEA/DMF N-TRITYLGLYCINE

5% TFA/5% TIS in CH₂Cl₂ BIOTIN/HBTU/HOBt/DIEA TFA/H₂0/PHENOL

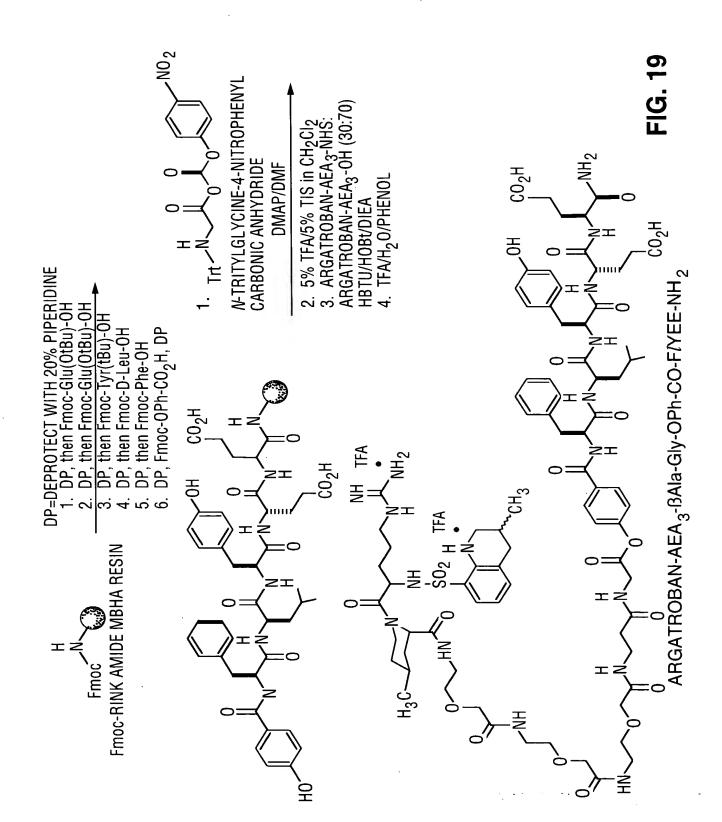
BIOTIN-Gly-OPh-CO-F/YEE-NH₂

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LC-BIOTIN ISOBUTYLCARBONIC ANHYDRIDE TEA/DMF 2. TFA/H₂O/PHENOL 土 **DP=DEPROTECT WITH 20% PIPERIDINE** c_0^2 H I Z DP, then Fmoc-Glu(0tBu)-0H DP, then Fmoc-Glu(0tBu)-0H DP, then Fmoc-Tyr(tBu)-OH DP, then Fmoc-D-Leu-OH DP, then Fmoc-Phe-OH DP, Fmoc-OPh-CO₂H, DP ZI 0: ა 4. ი. ი. Fmoc-RINK AMIDE MBHA RESIN

HN HN NH HN HN NH HN CO₂H CO₂H



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DP, then Fmoc-Glu(OtBu)-OF

SYNTHETIC SCHEMES

then Fmoc-Tyr(tBu)-(က

Fmoc-RINK AMIDE MBHA RESIN

Trt-AEA-0H

HBTU/HOBt/DIEA

0

C02H

FLUORESCEIN-THIOUREA-AEA $_3$ -Gly-OPh-CO-F $I\!Y$ EE-NH $_2$

I

DP, then Fmoc-Glu(OtBu)-OH

DP, then Fmoc-D-Leu-ÓH DP, then Fmoc-Phe-OH DP, Fmoc-OPh-CO₂H, DP 4. 7. 6

N-TRITYLGLYCINE-4-NITROPHENYL CARBONIC ANHYDRIDE

5% TFA/5% TIS in CH₂Cl₂ DMAP/DMF

5% TFA/5% TIS in CH₂Cl₂